

Claims:

1. A method of expressing a desired isoform of a gene product in a cell absent
5 undesired isoforms of a gene product, said method comprising:
 - (a) exposing a mammalian cell to at least one nucleic acid, said
nucleic acid being at least a partially double-stranded
ribonucleic acid and the double-stranded portion having at least
95% sequence identity to a common nucleic acid sequence
10 shared by two or more isoforms of said gene product; and
 - (b) introducing an expression vector encoding a desired isoform of
said gene product into said mammalian cell, said desired
isoform having a sequence comprising one or more mismatches
relative to said double-stranded portion of said nucleic acid,
15 operably linked to a promoter capable of driving expression of
said desired isoform in said cell.
2. The method of claim 1, wherein said common nucleic acid sequence is at
least 19 consecutive nucleotides in length.
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3. The method of claim 1 or 2, wherein said common nucleic acid sequence is
common to all endogenous isoforms of said gene product in said cell.
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4. The method of any one of claims 1 to 3, wherein the double-stranded portion
25 of said nucleic acid is 100% identical to said common nucleic acid sequence.
5. The method of any one of claims 1 to 4, wherein said nucleic acid is 19 to 25
nucleotides long.
- 30 6. The method of any one of claims 1 to 5, wherein said at least partially double-
stranded ribonucleic acid comprises a double-stranded portion of at least 19
nucleotides and at least one two-nucleotide single-stranded 3' overhang.

7. The method of any one of claims 1 to 6, wherein said desired isoform comprises a sequence comprising two or more mismatches relative to said double-stranded portion of said nucleic acid.

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8. The method of any of claims 1 to 7, wherein said expression vector encodes said desired isoform using at least one codon that differs from the endogenous sequence coding said desired isoform.

10 9. The method of claim 8, wherein said expression vector encodes said desired isoform using two codons that differ from the corresponding endogenous sequence coding said desired isoform.

15 10. The method of claim 8 or 9, wherein said desired isoform has an identical protein sequence to the corresponding endogenous isoform.

11. The method of any one of claims 1 to 10, wherein said desired isoform replaces a mutant isoform in the cell.

20 12. The method of claim 11, wherein said mutant isoform is oncogenic, apoptotic, tumor suppressive, inflammation inducive or suppressive, or angiogenic.

13. The method of any one of claims 1 to 12, further comprising determining the function of said desired isoform.

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14. The method of any one of claims 1 to 13, wherein said cell is a cancer cell.

15. The method of claim 14, wherein said cell is selected from the group consisting of HeLa (cervical cancer), PC3 (prostate cancer), MDA-MB-231 (breast cancer) and MCF-7.

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16. The method of any one of claims 1 to 15, wherein said desired isoform is transcribed under the control of an endogenous promoter.

17. The method of any one of claims 1 to 16, wherein said expression vector
5 comprises a constitutive promoter operably linked to said desired isoform.

18. The method of any one of claims 1 to 16, wherein said expression vector comprises an inducible promoter operably linked to said desired isoform.

10 19. The method of any one of claims 1 to 16, wherein said expression vector comprises a tissue-specific promoter operably linked to said desired isoform.

20. A kit comprising reagents expressing a desired isoform of a gene product in a cell absent undesired isoforms of a gene product, wherein said kit comprises
15 a nucleic acid being at least a partially double-stranded ribonucleic acid and the double-stranded portion having at least 95% sequence identity to a common nucleic acid sequence shared by two or more isoforms of said gene product; and an expression vector encoding a desired isoform of said gene product, said desired isoform having a sequence comprising one or more
20 mismatches relative to said double-stranded portion of said nucleic acid, operably linked to a promoter capable of driving expression of said desired isoform in said cell.

21. A mammalian cell exhibiting isoform-specific expression achieved by any of
25 the methods of claims 1-19.

22. A method for treating a disease comprising administering to a subject in need of treatment an effective amount of a nucleic acid being at least a partially double-stranded ribonucleic acid and the double-stranded portion having at
30 least 95% sequence identity to a common nucleic acid sequence shared by two or more isoforms of said gene product; and an expression vector

encoding a desired isoform of said gene product, said desired isoform having a sequence comprising one or more mismatches relative to said double-stranded portion of said nucleic acid, operably linked to a promoter capable of driving expression of said desired isoform in said cell.

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23. A method of assigning function to a desired isoform, said method comprising:

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a) exposing a mammalian cell to at least one nucleic acid, said nucleic acid being at least a partially double-stranded ribonucleic acid and the double-stranded portion having at least 95% sequence identity to a common nucleic acid sequence shared by two or more isoforms of said gene product;

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b) exposing said mammalian cell to an expression vector encoding a desired isoform of said gene product, said desired isoform having a sequence comprising one or more mismatches relative to said double-stranded portion of said nucleic acid, operably linked to a promoter capable of driving expression of said desired isoform in said cell;

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c) identifying a phenotype of said mammalian cell compared to when said desired isoform is absent, and
d) assigning said phenotype or function to said desired isoform.